(12) INTERNATIONAL APPLICATION PUBLISHED UNDER THE PATENT COOPERATION TREATY (PCT)

(19) World Intellectual Property Organization International Bureau



(43) International Publication Date 4 January 2001 (04.01.2001)

(10) International Publication Number WO 01/00882 A1

(51) International Patent Classification7: C12P 19/34, C07H 21/04

C12Q 1/68,

(21) International Application Number: PCT/US00/40266

(22) International Filing Date: 21 June 2000 (21.06.2000)

(25) Filing Language:

English

(26) Publication Language:

English

(30) Priority Data: 60/141,160

25 June 1999 (25.06.1999) US

(71) Applicant (for all designated States except US): THE TRUSTEES OF THE UNIVERSITY OF PENNSYL-VANIA [US/US]; Center for Technology Transfer, Suite

300, 3700 Market Street, Philadelphia, PA 19107 (US).

(72) Inventors: and

(75) Inventors/Applicants (for US only): EBERWINE, James, H. [US/US]; 3918 Henry Avenue, Philadelphia. PA 19129 (US). HEMBY, Scott, E. [US/US]; 450 Fourth Fairway Drive, Roswell, GA 30076 (US).

(74) Agents: LICATA, Jane, Massey et al.; Law Offices of Jane Massey Licata, 66 E. Main Street, Marlton, NJ 08053 (81) Designated States (national): AE, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CR, CU, CZ, DE, DK, DM, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW.

(84) Designated States (regional): ARIPO patent (GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW), Eurasian patent (AM, AZ, BY, KG, KZ, MD, RU, TJ, TM), European patent (AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE), OAPI patent (BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG).

Published:

- With international search report.
- Before the expiration of the time limit for amending the claims and to be republished in the event of receipt of amendments.

For two-letter codes and other abbreviations, refer to the "Guidance Notes on Codes and Abbreviations" appearing at the beginning of each regular issue of the PCT Gazette.

(54) Title: MOLECULAR CORRELATES OF SCHIZOPHRENIA AND METHODS OF DIAGNOSING SCHIZOPHRENIA VIA THESE MOLECULAR CORRELATES

(57) Abstract: Molecular correlates of genes and expressed sequence tags useful in the diagnosis and monitoring of treatment of patients suffering from schizophrenia are provided.

PATENT COOPERATION TREATY

M	Zermiren.
	OCT 2 6 2000
111	

From the INTERNATIONAL SEARCHING AUTHORITY

THOM the INTERNATIONAL DELIVERY	DCT	
To: JANE MASSEY LICATA LAW OFFICES OF JANE MASSEY LICATA 66 E. MAIN STREET	rc1	
MARLTON, NEW JERSEY 08053 Docket System	NOTIFICATION OF TRANSMITTAL OF THE INTERNATIONAL SEARCH REPORT OR THE DECLARATION	
Status Report Docket Book • 12/24/00 AMS	(PCT Rule 44.1)	
· 1:1124/00 W-	Date of Mailing (day/month/year) 24 OCT 2000	
i di anti di a		
Applicant's or agent's file reference PENN-0718	FOR FURTHER ACTION See paragraphs 1 and 4 below	
International application No.	International filing date (day/month/year) 21 JUNE 2000	
PCT/US00/40266	2170/12/2000	
Applicant The Trustees of the University of Pennsylvania		
1. X The applicant is hereby notified that the international	al search report has been established and is transmitted herewith.	
Till and amounts and statement under Artic		
The time limit for filing such amends	nents is normally 2 months from the date of transmittal of the or more details, see the notes on the accompanying sheet.	
Where? Directly to the International Bureau of 34, chemin des Colomb	WIPO	
1211 Geneva 20, Switze Facsimile No.: (41-22)	erland	
For more detailed instructions, see the notes of	n the accompanying sheet.	
2. The applicant is hereby notified that no internation Article 17(2)(a) to that effect is transmitted herewi	al search report will be established and that the declaration under th.	
3. With regard to the protest against payment of (a	n) additional fee(s) under Rule 40.2, the applicant is notified that:	
the protest together with the decision thereon applicant's request to forward the texts of b	has been transmitted to the International Bureau together with the oth the protest and the decision thereon to the designated Offices.	
no decision has been made yet on the prote	st; the applicant will be notified as soon as a decision is made.	
4. Further action(s): The applicant is reminded of the following:		
Shortly after 18 months from the priority date, the international application will be published by the International Bureau. If the applicant wishes to avoid or postpone publication, a notice of withdrawal of the international application, or of the priority claim, must reach the International Bureau as provided in rules 90 bis 1 and 90 bis 3, respectively, before the completion of the technical preparations for international publication.		
Within 19 months from the priority date, a demand for international preliminary examination must be filed if the applicant wishes to postpone the entry into the national phase until 30 months from the priority date (in some Offices even later).		
True to an all from the priority date the applicant	must perform the prescribed acts for entry into the national phase ected in the demand or in a later election within 19 months from the	
C. 704 / TO	Authorized of the last	
Name and mailing address of the ISA/US Commissioner of Patents and Trademarks	JEHANNE SODATA	
Box PCT Washington, D.C. 20231	JEHAMIC SOPATA	
Facsimile No. (703) 305-3230	Telephone No. (703) 308-1235	

Form PCT/ISA/220 (July 1998)★

(See notes on accompanying sheet)

PATENT COOPERATION TREATY

......

From the INTERNATIONAL SEARCHING AUTHORITY			
To: JANE MASSEY LICATA LAW OFFICES OF JANE MASSEY LICATA 66 E. MAIN STREET	PCT		
MARLTON, NEW JERSEY 08053	NOTIFICATION OF TRANSMITTAL OF THE INTERNATIONAL SEARCH REPORT OR THE DECLARATION		
•	(PCT Rule 44.1)		
	Date of Mailing (day/month/year) 24 OCT 2000		
Applicant's or agent's file reference PENN-0718	FOR FURTHER ACTION See paragraphs 1 and 4 below		
International application No. PCT/US00/40266	International filing date (day/month/year) 21 JUNE 2000		
Applicant The Trustees of the University of Pennsylvania			
1. X The applicant is hereby notified that the international	al search report has been established and is transmitted herewith.		
	the claims of the international application (see Rule 40).		
When? The time limit for filing such amendn international search report; however, fo	nents is normally 2 months from the date of transmittal of the or more details, see the notes on the accompanying sheet.		
Where? Directly to the International Bureau of 34, chemin des Colomb 1211 Geneva 20, Switze Facsimile No.: (41-22)	ettes erland		
For more detailed instructions, see the notes on the accompanying sheet.			
2. The applicant is hereby notified that no internation Article 17(2)(a) to that effect is transmitted herewise	al search report will be established and that the declaration under th.		
	n) additional fee(s) under Rule 40.2, the applicant is notified that:		
applicant's request to forward the texts of be	has been transmitted to the International Bureau together with the oth the protest and the decision thereon to the designated Offices.		
no decision has been made yet on the protest; the applicant will be notified as soon as a decision is made.			
4. Further action(s): The applicant is reminded of the following:			
Shortly after 18 months from the priority date, the international application will be published by the International Bureau. If the applicant wishes to avoid or postpone publication, a notice of withdrawal of the international application, or of the priority claim, must reach the International Bureau as provided in rules 90 bis 1 and 90 bis 3, respectively, before the completion of the technical preparations for international publication.			
Within 19 months from the priority date, a demand for international preliminary examination must be filed if the applicant wishes to postpone the entry into the national phase until 30 months from the priority date (in some Offices even later).			
Within 20 months from the priority date, the applicant must perform the prescribed acts for entry into the national phase before all designated Offices which have not been elected in the demand or in a later election within 19 months from the priority date or could not be elected because they are not bound by Chapter			
	AllA		
Name and mailing address of the ISA/US	Authorized office		
Commissioner of Patents and Trademarks	JEHAMNE SODAYA		
Box PCT Washington, D.C. 20231			
Facsimile No. (703) 305-3230	Telephone No. (703) 308-1235		

Form PCT/ISA/220 (July 1998)*

(See notes on accompanying sheet)

PATENT COOPERATION TREATY

PCT

INTERNATIONAL SEARCH REPORT

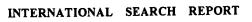
(PCT Article 18 and Rules 43 and 44)

Applicant's or agent's file reference PENN-0718	FOR FURTHER ACTION	see Notification of (Form PCT/ISA/220	Transmittal of Internal) as well as, where app	olicable, item 3 below.
International application No.	International filing dat	e (day/month/year)	(Earliest) Priority Da	te (day/month/year)
PCT/US00/40266	21 JUNE 2000		25 JUNE 1999	
Applicant The Trustees of the University of Pen	nsylvania			
This international search report has be according to Article 18. A copy is bei	\int	mational Bulcau.	othority and is transm	itted to the applicant
This international search report consis	ts of a total of <u>#</u> shee	ts.		
X It is also accompanied by a	copy of each prior art do	cument cited in this	report.	
With regard to the language, language in which it was filed the international search was Authority (Rule 23.1(b)). With regard to any nucleotide was carried out on the basis of the language.	I, unless otherwise indicate is carried out on the basis and/or amino acid seque	of a translation of t	he international appli	cation furnished to this
contained in the internatio		form.		
filed together with the inte	rnational application in c	omputer readable for	m.	
furnished subsequently to				
furnished subsequently to	this Authority in compute	er readable form.		
the statement that the substinternational application a the statement that the inforfurnished.	equently furnished written is filed has been furnishe mation recorded in comput	n sequence listing do d. er readable form is id		
2. Certain claims were fou		OX 1).		
3. Unity of invention is lac	King (See Box 11).			
4. With regard to the title, X the text is approved as so	ibmitted by the applicant.			
the text is approved as si the text has been establis				
	ubmitted by the applicant hed, according to Rule 30 ay, within one month from	R 2/h) by this Autho	rity as it appears in of this international	
search report, submit co	mments to this Authority	•		
6. The figure of the drawings to b		ract is rigure inc		None of the figures.
as suggested by the app			L	None of the figures.
because the applicant fa	iled to suggest a figure.			

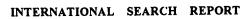
International application No. PCT/US00/40266

A. CLASSIFICATION OF SUBJECT MATTER			
	C12Q 1/68; C12P 19/34; CO7H 21/04		
According to	US CL: 435/6, 91.1; 536/23.1 According to International Patent Classification (IPC) or to both national classification and IPC		
B. FIELI	DS SEARCHED		
Minimum do	cumentation searched (classification system followed	by classification symbols)	
U.S. : 4	35/6, 91.1; 536/23.1		
	on searched other than minimum documentation to the e	week that such documents are included	in the fields searched
Documentati	on searched other than minimum documentation to the e	xtent utat such documents are incided	In the Herat controller
	•		
Electronic da	ata base consulted during the international search (nam	ne of data base and, where practicable	e, search terms used)
	Extra Sheet.		
C. DOCI	UMENTS CONSIDERED TO BE RELEVANT		
· · · · · · · · · · · · · · · · · · ·	Citation of document, with indication, where app	ropriate of the relevant nassages	Relevant to claim No.
Category*	Citation of document, with indication, where app.	Topriate, of the relevant passages	
$ _{\mathbf{Y}}$	EASTWOOD S. L. et al. Hippocam	pal and Cortical Growth-	1-3
	Associated Protein-43 Messenger		
	Neuroscience. 1998. Vol 86. pages 437	-448, especially abstract.	
	GOVOYOUR R. Commercian of MNDAE	01 GluD1 GluB7 and KA1	1
X	SOKOLOV B.P. Expression of MNDAR Glutamate Receptor mRNAs is Decre	ased in Frontal Cortex of	1 - 1
Y	'Neuroleptic-Free' Schizophrenics: Ev	idence on Reversible Up-	2-3
1	Regulation by Typical Neuroleptics.	Journal of Neurochemistry.	
	1998. Vol 71. pages 2454-2464, especia	ally abstract.	
	'		
		•	
X Furt	her documents are listed in the continuation of Box C	. See patent family annex.	
• s _i	pecial categories of cited documents:	"T" later document published after the indate and not in conflict with the ap	plication but cited to understand
"A" do	ocument defining the general state of the art which is not considered be of particular relevance	the principle or theory underlying t	he invention
	arlier document published on or after the international filing date	"X" document of particular relevance, considered novel or cannot be consi	the claimed invention cannot be dered to involve an inventive step
L document which may throw doubts on priority claim(s) or which is when the document is taken alone			
•5	pecial reason (as specified)	 Y's document of particular relevance; considered to involve an inventi combined with one or more other s 	ve step when the document is
m	ocument referring to an oral disclosure, use, exhibition or other seams	being obvious to a person skilled i	n the art
tì	ocument published prior to the international filing date but later than he priority date claimed	*& document member of the same pat	
Date of the	e actual completion of the international search	Date of mailing of the international	search report
27 SEPTEMBER 2000 24 027 2000			
Name and mailing address of the ISA/US Authorized officer V 000			
Commissioner of Patents and Trademarks Box PCT JEHANNE SOUAYA			
Washingt	on, D.C. 20231 No. (703) 305-3230	Telephone No. (703) 308-1235	//
I Maccimile	1945 1 (14.5) 34 (2.4.3)		11

Form PCT/ISA/210 (second sheet) (July 1998)*



C (Continua	tion). DOCUMENTS CONSIDERED TO BE RELEVANT		
Category*	Citation of document, with indication, where appropriate, of the relev	ant passages	Relevant to claim No.
X Y	GLANTZ L.A. et al. Reduction of Synaptophysin Immunoreactivity in the Prefrontal Cortex of Subjects of Schizophrenia. Archives of General Psychiatry. October 54. pages 943-952, especially abstract.	with .	1 2-3
	·		
	,		1
	·		
į			



International application No.

PCT/US00/40266

B. FIELDS SEARCHED Electronic data bases consulted (Name of data base and where practicable terms used):	
STN, Medline, Caplus, Uspatful search terms: schizophrenia, diagnose, microarray, adaptin, synaptic vessicle transporter, synaptotagmin 1, gynaptotagmin IV, Gaba transporter 1	
gynaptolaginin 17, Guou transporter 1	
•	
·	
·	
	_
	•

B. FIELDS SEARCHED Electronic data bases consulted (Name of data base and where practicable terms used):	
STN, Medline, Caplus, Uspatful search terms: schizophrenia, diagnose, microarray, adaptin, synaptic vessicle transporter, synaptotagmin 1, gynaptotagmin IV, Gaba transporter 1	

International application No. PCT/US00/40266

A. CLASSIFICATION OF SUBJECT MATTER IPC(7) :C12Q 1/68; C12P 19/34; C07H 21/04 US CL :435/6, 91.1; 536/23.1 According to International Patent Classification (IPC) or to both national classification and IPC			
B. FIELDS SEARCHED			
Minimum documentation searched (classification system follo-	wed by classification symbols)		
U.S. : 435/6, 91.1; 536/23.1	,, ,		
Documentation searched other than minimum documentation to	the extent that such documents are included in the fields searched		
Electronic data base consulted during the international search Please See Extra Sheet.	(name of data base and, where practicable, search terms used)		
C. DOCUMENTS CONSIDERED TO BE RELEVANT			
Category* Citation of document, with indication, where	appropriate, of the relevant passages Relevant to claim No.		
Y EASTWOOD S. L. et al. Hippoc Associated Protein-43 Messenger Neuroscience. 1998. Vol 86. pages 4	RNA in Schizophrenia		
SOKOLOV B.P. Expression of MND. Glutamate Receptor mRNAs is Dec 'Neuroleptic-Free' Schizophrenics: Regulation by Typical Neuroleptics. 1998. Vol 71. pages 2454-2464, espe	AR1, GluR1, GluR7, and KA1 creased in Frontal Cortex of Evidence on Reversible Up- Journal of Neurochemistry		
Special categories of cited documents: A** document defining the general state of the art which is not considered to be of particular relevance to be of particular relevance document which may throw doubts on priority claim(s) or which is cited to establish the publication date of another citation or other special reason (as specified) B** document referring to an oral disclosure, use, exhibition or other means C** document published prior to the international filing date but later than the priority date claimed C** document published prior to the international filing date but later than the priority date claimed C** document published prior to the international filing date but later than the priority date claimed C** document published prior to the international search C** document published prior to the international filing date but later than the priority date claimed C** document published prior to the international filing date but later than the priority date claimed C** document published prior to the international filing date but later than the priority date claimed C** document published prior to the international filing date but later than the priority date claimed C** document published prior to the international filing date but later than the priority date claimed C** document published after the international filing date or priority date and not in conflict with the application but cited to understand the priority date and not in conflict with the application but cited to understand the priority date and not in conflict with the application but cited to understand the priority date and not in conflict with the application but cited to understand the priority date and not in conflict with the application but cited to understand the priority date and not in conflict with the application but cited to understand the prio			
Name and mailing address of the ISA/US Commissioner of Patents and Trademarks Box PCT Washington, D.C. 20231 Facsimile No. (703) 305-3230 Authorized officer JEHANNE SOUAYA Telephone No. (703) 308-1235			

Form PCT/ISA/210 (second sheet) (July 1998) *

		101/0300/4020	· ·
C (Continua	tion). DOCUMENTS CONSIDERED TO BE RELEVANT		
Category*	Citation of document, with indication, where appropriate, of the relev	ant passages	Relevant to claim No
X Y	GLANTZ L.A. et al. Reduction of Synaptophysin Immunoreactivity in the Prefrontal Cortex of Subjects v Schizophrenia. Archives of General Psychiatry. October 54. pages 943-952, especially abstract.	with 1997. Vol	1 2-3

B. FIELDS SEARCHED Electronic data bases consulted (Name of data base and where practicable terms used):
STN, Medline, Caplus, Uspatful search terms: schizophrenia, diagnose, microarray, adaptin, synaptic vessicle transporter, synaptotagmin 1, gynaptotagmin IV, Gaba transporter 1
,



PCT

NOTIFICATION OF ELECTION

(PCT Rule 61.2)

To:

Commissioner
US Department of Commerce
United States Patent and Trademark
Office, PCT
2011 South Clark Place Room
CP2/5C24
Arlington, VA 22202

Date of mailing (day/month/year) 22 August 2001 (22.08.01)	ETATS-UNIS D'AMERIQUE in its capacity as elected Office
International application No. PCT/US00/40266	Applicant's or agent's file reference PENN-0718
International filing date (day/month/year) 21 June 2000 (21.06.00)	Priority date (day/month/year) 25 June 1999 (25.06.99)
Applicant	
FRERWINE lames H ot al	

1.	The designated Office is hereby notified of its election made:
	X in the demand filed with the International Preliminary Examining Authority on:
	12 January 2001 (12.01.01)
	in a notice effecting later election filed with the International Bureau on:
2.	The election X was
	was not
	made before the expiration of 19 months from the priority date or, where Rule 32 applies, within the time limit under Rule 32.2(b).

The International Bureau of WIPO 34, chemin des Colombettes 1211 Geneva 20, Switzerland Authorized officer

Eugénia SANTOS (Fax 338.87.40)

Telephone No.: (41-22) 338.83.38

Facsimile No.: (41-22) 740.14.35